## AMENDMENT TO THE CLAIMS

- 1. (Withdrawn) A method of producing an Isotope Map for data from a mass spectometric injection of a biological sample comprising:
- a) performing Noise reduction and Centroiding with a Noise Reduction Module on said data from a mass spectometric injection of a biological sample, and
- b) generating an Isotope Map from the noise reduced and centroided data, thereby producing an Isotope Map.
- 2. (Withdrawn) The method of claim 1, wherein said biological sample is comprised of unlabeled biomolecules.
- 3. (Withdrawn) The method of claim 1, wherein said biological sample is comprised of underivatized biomolecules.
- 4. (Withdrawn) The method of claim 1, wherein said biological sample is comprised of biomolecules that are both unlabeled and underivatized.
- 5. (Withdrawn) The method of claim 1, wherein said biological sample is comprised of cleaved biomolecules.
- 6. (Withdrawn) The method of claim 5, wherein said biomolecules are cleaved with an enzyme.
- 7. (Withdrawn) The method of claim 6, wherein said enzyme is trypsin.

- 8. (Withdrawn) A method for producing a Peptide Map from an Isotope Map for data from a mass spectometric injection of a biological sample comprising:
- a) performing Peptide Detection on an Isotope Map with a Peptide Detection Module, and
- b) generating a Peptide Map from the results of the peptide detection, thereby producing a Peptide Map.
- 9. (Currently Amended) A method for determining the functions to align Peptide Maps from a pair of Peptide Maps, each representing a mass spectrometric injection of a biological sample, comprising:
- a) performing Peptide Map Alignment on a pair of Peptide Maps with a Peptide Map Alignment Module, and
- b) determining a column offset between the pair of injections, and
- c) determining a retention time transformation function between the pair of injections.
- 10. (Original) A method for Alignment of Peptide Maps from a pair of Peptide Maps, each representing a mass spectrometric injection of a biological sample, comprising performing Peptide Map Alignment on a pair of Peptide Maps with a Peptide Map Alignment Module, and thereby producing Aligned Peptide Maps.
- (Currently Amended) [[A]] The method of claim 10, further comprising determining differentially and uniquely expressed biomolecules from a pair of mass spectrometric injections of biological samples, comprising:
- a) producing Isotope Maps for the samples;
- b) producing Peptide Maps from the Isotope Maps;
- c) aligning the Peptide Maps, and
- d) determining the differences differential intensities of biomolecules between the aligned

maps, thereby determining differentially and uniquely expressed biomolecules from the pair of injections.

- (Currently Amended) [[A]] The method of claim 10, further comprising matching biomolecules a precursor from which an LC-MS injection to LC-MS-MS fragmentation spectra spectrum was obtained to the precursor in one of said Peptide Maps-from comprising:
- a) producing Isotope Maps for the injections;
- b) producing Peptide Maps from the Isotope Maps;
- c) aligning the Peptide Maps, and
- d) determining the matching biomolecules.
- 13. (Withdrawn) A computer implemented method of producing an Isotope Map for data from a mass spectometric injection of a biological sample comprising:
- a) inputting data from a mass spectometric injection of a biological sample, and performing Noise reduction and Centroiding with a Noise Reduction Module on said, and
- b) generating an Isotope Map from the noise reduced and centroided data, thereby producing an Isotope Map.
- 14. (Withdrawn) The computer implemented method of claim 13, wherein said biological sample is comprised of unlabeled biomolecules.
- 15. (Withdrawn) The computer implemented method of claim 13, wherein said biological sample is comprised of underivatized biomolecules.
- 16. (Withdrawn) The computer implemented method of claim 13, wherein said

biological sample is comprised of biomolecules that are both unlabeled and underivatized.

- 17. (Withdrawn) The computer implemented method of claim 13, wherein said biological sample is comprised of cleaved biomolecules.
- 18. (Withdrawn) The computer implemented method of claim 17, wherein said biomolecules are cleaved with an enzyme.
- 19. (Withdrawn) The computer implemented method of claim 18, wherein said enzyme is trypsin.
- 20. (Withdrawn) A computer implemented method for producing a Peptide Map from an Isotope Map for data from a mass spectometric injection of a biological sample comprising:
- a) inputting an Isotope Map;
- b) performing Peptide Detection on said Isotope Map with a Peptide Detection Module, and
- c) generating a Peptide Map from the results of the peptide detection, thereby producing a Peptide Map.
- 21. (Original) A computer implemented method for determining the functions to align Peptide Maps from a pair of Peptide Maps, each representing a mass spectrometric injection of a biological sample, comprising:
- a) inputting a pair of Peptide Maps;
- b) performing Peptide Map Alignment on a pair of Peptide Maps with a Peptide Map Alignment Module;
- c) determining a column offset between the pair of injections, and

- d) determining a retention time transformation function between the pair of injections.
- 22. (Original) A computer implemented method for Alignment of Peptide Maps from a pair of Peptide Maps, each representing a mass spectrometric injection of a biological sample, comprising inputting a pair of Peptide Maps, performing Peptide Map Alignment on said pair of Peptide Maps with a Peptide Map Alignment Module, and thereby producing Aligned Peptide Maps.
- 23. (Currently Amended) [[A]] <u>The</u> computer implemented method of <u>claim 22</u>, <u>further comprising determining differentially and uniquely expressed biomolecules from a pair of mass spectrometric injections of biological samples, comprising:</u>
- a) inputting data from said injections;
- b) producing Isotope Maps for the samples;
- c) producing Peptide Maps from the Isotope Maps;
- d) aligning the Peptide Maps, and e) determining the differences differential intensities of biomolecules between the aligned maps, thereby determining differentially and uniquely expressed biomolecules from the pair of injections.
- 24. (Currently Amended) [[A]] <u>The</u> computer implemented method of <u>claim 22</u>, <u>further comprising</u> matching <u>biomolecules</u> <u>a precursor</u> from <u>which</u> an <u>LC-MS injection to</u> LC-MS-MS fragmentation <u>spectra</u> <u>spectrum was obtained to the precursor in one of said</u> <u>Peptide Maps from comprising:</u>
- a) inputting data from said injections;
- b) producing Isotope Maps for the injections;
- c) producing Peptide Maps from the Isotope Maps;
- d) aligning the Peptide Maps, and
- e) determining the matching biomolecules.

25. (Withdrawn) A computer-readable memory having stored thereon a program for producing an Isotope

Map for data from a mass spectometric injection of a biological sample comprising:

- a) computer code that receives as input data from a mass spectometric injection of a biological sample, and performing Noise reduction and Centroiding with a Noise Reduction Module on said, and
- b) and computer code that generates an Isotope Map from the noise reduced and centroided data, thereby producing an Isotope Map.
- 26. (Withdrawn) The computer-readable memory of claim 25, wherein said biological sample is comprised of unlabeled biomolecules.
- 27. (Withdrawn) The computer-readable memory of claim 25, wherein said biological sample is comprised of underivatized biomolecules.
- 28. (Withdrawn) The computer-readable memory of claim 25, wherein said biological sample is comprised of biomolecules that are both unlabeled and underivatized.
- 29. (Withdrawn) The computer-readable memory of claim 25, wherein said biological sample is comprised of cleaved biomolecules.
- 30. (Withdrawn) The computer-readable memory of claim 29, wherein said biomolecules are cleaved with an enzyme.
- 31. (Withdrawn) The computer-readable memory of claim 30, wherein said enzyme is trypsin.

- 32. (Withdrawn) A computer-readable memory having stored thereon a program for producing a Peptide Map from an Isotope Map for data from a mass spectometric injection of a biological sample comprising:
- a) computer code that receives as input an Isotope Map;
- b) computer code that performs Peptide Detection on said Isotope Map with a Peptide Detection Module, and
- c) computer code that generates a Peptide Map from the results of the peptide detection, thereby producing a Peptide Map.
- 33. (Original) A computer-readable memory having stored thereon a program for determining the functions to align Peptide Maps from a pair of Peptide Maps, each representing a mass spectrometric injection of a biological sample, comprising:
- a) computer code that receives as input a pair of Peptide Maps;
- b) computer code that performs Peptide Map Alignment on a pair of Peptide Maps with a Peptide Map Alignment Module;
- c) computer code that determines a column offset between the pair of injections, and
- d) computer code that determines a retention time transformation function between the pair of injections.
- 34. (Original) A computer-readable memory having stored thereon a program for Alignment of Peptide Maps from a pair of Peptide Maps, each representing a mass spectrometric injection of a biological sample, comprising computer code that receives as input a pair of Peptide Maps, and computer code that performs Peptide Map Alignment on said pair of Peptide Maps with a Peptide Map Alignment Module, and thereby produces Aligned Peptide Maps.

- 35. (Currently Amended) [[A]] <u>The computer-readable memory of claim 34, wherein having stored thereon a the program further comprises for determining differentially and uniquely expressed biomolecules from a pair of mass spectrometric injections of biological samples, comprising:</u>
- a) computer code that receives as input data from said injections;
- b) computer code that produces Isotope Maps for the samples;
- c) computer code that produces Peptide Maps from the Isotope Maps;
- d) computer code that aligns the Peptide Maps, and
- e) computer code that determines the <u>differences</u> <u>differential intensities of biomolecules</u> between the aligned maps, <u>thereby determining differentially and uniquely expressed</u> biomolecules from the pair of injections.
- 36. (Currently Amended) [[A]] <u>The computer-readable memory of claim 34, wherein having stored thereon a the program further comprises computer code for matching biomolecules a precursor from which an LC-MS injection to LC-MS-MS fragmentation spectra spectrum was obtained to the precursor in one of said Peptide Maps from comprising:</u>
- a) computer code that receives as input data from said injections;
- b) computer code that produces Isotope Maps for the injections;
- c) computer code that produces Peptide Maps from the Isotope Maps;
- d) computer code that aligns the Peptide Maps, and
- e) computer code that determines the matching biomolecules.
- 37. (Withdrawn) A computer system for producing an Isotope Map for data from a mass spectometric injection of a biological sample comprising a processor and a memory

- coupled to said processor, said memory encoding one or more programs, said one or more programs causing said processor to perform a method comprising:
- a) computer code that receives as input data from a mass spectometric injection of a biological sample, and performing Noise reduction and Centroiding with a Noise Reduction Module on said, and
- b) computer code that generates an Isotope Map from the noise reduced and centroided data, thereby producing an Isotope Map.
- 38. (Withdrawn) The computer system of claim 37, wherein said biological sample is comprised of unlabeled biomolecules.
- 39. (Withdrawn) The computer system of claim 37, wherein said biological sample is comprised of underivatized biomolecules.
- 40. (Withdrawn) The computer system of claim 37, wherein said biological sample is comprised of biomolecules that are both unlabeled and underivatized.
- 41. (Withdrawn) The computer system of claim 37, wherein said biological sample is comprised of cleaved biomolecules.
- 42. (Withdrawn) The computer system of claim 41, wherein said biomolecules are cleaved with an enzyme.
- 43. (Withdrawn) The computer system of claim 42, wherein said enzyme is trypsin.
- 44. (Withdrawn) A computer system for producing a Peptide Map from an Isotope Map for data from a mass spectometric injection of a biological sample comprising a

processor and a memory coupled to said processor, said memory encoding one or more programs, said one or more

programs causing said processor to perform a method comprising:

- a) computer code that receives as input an Isotope Map;
- b) computer code that performs Peptide Detection on said Isotope Map with a Peptide Detection Module, and
- c) computer code that generates a Peptide Map from the results of the peptide detection, thereby producing a Peptide Map.
- 45. (Currently Amended) A computer system for determining the functions to align Peptide Maps from a pair of Peptide Maps, each representing a mass spectrometric injection of a biological sample, comprising a processor and a memory coupled to said processor, said memory encoding one or more programs, said one or more programs causing said processor to perform a method comprising:
- a) computer code that receives receiving as input a pair of Peptide Maps;
- b) computer code that performs performing Peptide Map Alignment on a pair of Peptide Maps with a Peptide Map Alignment Module;
- c) computer code that determines determining a column offset between the pair of injections, and
- d) computer code that determines determining a retention time transformation function between the pair of injections.
- 46. (Currently Amended) A computer system for Alignment of Peptide Maps from a pair of Peptide Maps, each representing a mass spectrometric injection of a biological sample, comprising a processor and a memory coupled to said processor, said memory encoding one or more programs, said one or more programs causing said processor to perform a method comprising computer code that receives receiving as input a pair of

Peptide Maps, and computer code that performs performing Peptide Map Alignment on said pair of Peptide Maps with a Peptide Map Alignment Module, and thereby produces producing Aligned Peptide Maps.

- (Currently Amended) [[A]] The computer system of claim 46, wherein said 47. method performed by said processor further comprises for determining differentially and uniquely expressed biomolecules from a pair of mass spectrometric injections of biological samples, comprising a processor and a memory coupled to said processor, said memory encoding one or more programs, said one or more programs causing said processor to perform a method comprising:
- a) computer code that receives as input data from said injections;
- b) computer code that produces Isotope Maps for the samples;
- c) computer code that produces Peptide Maps from the Isotope Maps;
- d) computer code that aligns the Peptide Maps, and
- e) computer code that determines determining the differences differential intensities of biomolecules between the aligned maps, thereby determining differentially and uniquely expressed biomolecules from the pair of injections.
- 48. (Currently Amended) [[A]] The computer system of claim 46, wherein said method performed by said processor further comprises for matching biomolecules a precursor from which an LC-MS injection to LC-MS-MS fragmentation spectra spectrum was obtained to the precursor in one of said Peptide Maps from comprising a processor and a memory coupled to said processor, said memory encoding one or more programs, said one or more programs causing said processor to perform a method comprising: a) computer code that receives as input data from said injections;
- b) computer code that produces Isotope Maps for the injections;
- c) computer code that produces Peptide Maps from the Isotope Maps;

- d) computer code that aligns the Peptide Maps, and
- e) computer code that determines the matching biomolecules.
- 49. (Withdrawn) A method for displaying information on an Isotope Map for data from a mass spectometric injection of a biological sample comprising:
- a) inputting data from a mass spectometric injection of a biological sample;
- b) performing Noise reduction and Centroiding with a Noise Reduction Module on said data;
- c) generating an Isotope Map from the noise reduced and centroided data, thereby producing an Isotope Map, and
- d) displaying information on said Isotope Map to a user.
- 50. (Withdrawn) The method of claim 49, wherein said biological sample is comprised of unlabeled biomolecules.
- 51 (Withdrawn) The method of claim 49, wherein said biological sample is comprised of underivatized biomolecules.
- 52. (Withdrawn) The method of claim 49, wherein said biological sample is comprised of biomolecules that are both unlabeled and underivatized.
- 53. (Withdrawn) The method of claim 49, wherein said biological sample is comprised of cleaved biomolecules.
- 54. (Withdrawn) The method of claim 53, wherein said biomolecules are cleaved with an enzyme.

- 55. (Withdrawn) The method of claim 54, wherein said enzyme is trypsin.
- 56. (Withdrawn) A method for displaying information on a Peptide Map produced from an Isotope Map for data from a mass spectometric injection of a biological sample comprising:
- a) inputting an Isotope Map;
- b) performing Peptide Detection on said Isotope Map with a Peptide Detection Module;
- c) generating a Peptide Map from the results of the peptide detection, thereby producing a Peptide Map, and
- d) displaying information on said Peptide Map to a user.
- 57. (Original) A method for displaying information on the functions to align Peptide Maps from a pair of Peptide Maps, each representing a mass spectrometric injection of a biological sample, comprising:
- a) inputting a pair of Peptide Maps;
- b) performing Peptide Map Alignment on a pair of Peptide Maps with a Peptide Map Alignment Module;
- c) determining a column offset between the pair of injections;
- d) determining a retention time transformation function between the pair of injections, and
- e) displaying information on said functions to a user.
- 58. (Currently Amended) A method for displaying information on Alignment of Peptide Maps from a pair of Peptide Maps, each representing a mass spectrometric injection of a biological sample, comprising inputting a pair of Peptide Maps, and performing Peptide Map Alignment on said pair of Peptide Maps with a Peptide Map Alignment Module, thereby producing Aligned Peptide Maps, and displaying information

on said Aligned Peptide Maps to a user.

- 59. (Currently Amended) [[A]] The method of claim 58, further comprising for displaying information on differentially and uniquely expressed biomolecules from a pair of mass spectrometric injections of biological samples, comprising:
- a) inputting data from said injections;
- b) producing Isotope Maps for the samples;
- c) producing Peptide Maps from the Isotope Maps;
- d) aligning the Peptide Maps;
- e) determining the <u>differences</u> <u>differential intensities of biomolecules</u> between the aligned maps, <u>thereby determining differentially and uniquely expressed biomolecules from the pair of injections</u>, and <u>f</u>) displaying information on said <u>differences</u> <u>differential intensities</u> to <u>a the</u> user.
- 60. (Currently Amended) [[A]] The method of claim 58, further comprising for displaying information on matching biomolecules a precursor from which an LC-MS injection to LC-MS-MS fragmentation spectra spectrum was obtained to the precursor in one of said Peptide Maps from comprising:
- a) inputting data from said injections;
- b) producing Isotope Maps for the injections;
- c) producing Peptide Maps from the Isotope Maps;
- d) aligning the Peptide Maps, and
- e) determining the matching biomolecules,
- f) and displaying information on said matching biomolecules to a user.